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INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

<b>(51) International Patent Classification <sup>6</sup> :</b> <b>C07F 9/09</b>	<b>A1</b>	<b>(11) International Publication Number:</b> <b>WO 99/35150</b> <b>(43) International Publication Date:</b> 15 July 1999 (15.07.99)
<b>(21) International Application Number:</b> PCT/US99/00419 <b>(22) International Filing Date:</b> 8 January 1999 (08.01.99)  <b>(30) Priority Data:</b> 60/071,070 9 January 1998 (09.01.98) US 60/111,531 9 December 1998 (09.12.98) US  <b>(71) Applicant (for all designated States except US):</b> ARIZONA BOARD OF REGENTS, a body corporate, acting on behalf of ARIZONA STATE UNIVERSITY [US/US]; Tempe, AZ 85287 (US).  <b>(72) Inventors; and</b> <b>(75) Inventors/Applicants (for US only):</b> PETTIT, George, R. [US/US]; 6232 Bret Hills Drive, Paradise Valley, AZ 85253 (US). RHODES, Monte, R. [US/US]; 8115 Copper Creek Drive, Louisville, KY 40222 (US).  <b>(74) Agent:</b> MYBECK, Richard, R.; Suite 10, 8010 East Morgan Trail, Scottsdale, AZ 85258 (US).		<b>(81) Designated States:</b> CA, JP, US, European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).  <b>Published</b> <i>With international search report.</i> <i>With amended claims.</i>
<b>(54) Title:</b> SYNTHESIS OF COMBRETASTATIN A-4 PRODRUGS AND TRANS-ISOMERS THEREOF  <b>(57) Abstract</b>  Combretastatin A-4 has been previously selected for pre-clinical development as antineoplastic agents. However, it is essentially insoluble in water. New water soluble derivatives of combretastatin A-4 and its qualified analogs have been discovered and synthesized through a multistage process using other derivatives of combretastatin A-4 as intermediates. These water soluble derivatives are herein denominated as "Combretastatin A-4 Prodrugs".		